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(54) Title: NEW METHOD

(57) Abstract: The invention provides methods of screening test compounds for their ability to reduce retention of atherogenic lipoproteins in atherogenesis by measuring their activity as a modulator of the binding affinity of CEL (Carboxyl Ester Lipase) to various receptors. The invention further provides methods for the use of active modulators of the binding affinity of CEL to receptors for the prevention and treatment of atherosclerosis.

## A. CLASSIFICATION OF SUBJECT MATTER

IPC 7 G01N33/68 G01N33/92 A61K31/00 G01N33/573

According to International Patent Classification (IPC) or to both national classification and IPC

## B. FIELDS SEARCHED

Minimum documentation searched (classification system followed by classification symbols)

IPC 7 G01N A61K

Documentation searched other than minimum documentation to the extent that such documents are included in the fields searched

Electronic data base consulted during the International search (name of data base and, where practical, search terms used)

EPO-Internal, BIOSIS, EMBASE, COMPENDEX, PAJ, WPI Data, Sequence Search

## C. DOCUMENTS CONSIDERED TO BE RELEVANT

Category *	Citation of document, with indication, where appropriate, of the relevant passages	Relevant to claim No.
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X	<p>EP 0 640 620 A (LANGE, LOUIS G., III; BOSNER, MATTHEW S) 1 March 1995 (1995-03-01) the whole document in particular: abstract claims 3-5 column 1, line 18 - line 26 column 1, line 43 - column 2, line 9 column 4, line 45 - line 58 -----</p>	1-9
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☒ Further documents are listed in the continuation of box C.☒ Patent family members are listed in annex.

## \* Special categories of cited documents:

- "A" document defining the general state of the art which is not considered to be of particular relevance
- "E" earlier document but published on or after the international filing date
- "L" document which may throw doubts on priority claim(s) or which is cited to establish the publication date of another citation or other special reason (as specified)
- "O" document referring to an oral disclosure, use, exhibition or other means
- "P" document published prior to the international filing date but later than the priority date claimed

- "T" later document published after the international filing date or priority date and not in conflict with the application but cited to understand the principle or theory underlying the invention
- "X" document of particular relevance; the claimed invention cannot be considered novel or cannot be considered to involve an inventive step when the document is taken alone
- "Y" document of particular relevance; the claimed invention cannot be considered to involve an inventive step when the document is combined with one or more other such documents, such combination being obvious to a person skilled in the art.
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Date of the actual completion of the international search

16 September 2005

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## C.(Continuation) DOCUMENTS CONSIDERED TO BE RELEVANT

Category *	Citation of document, with indication, where appropriate, of the relevant passages	Relevant to claim No.
X	<p>BENGTSOON SARA H M ET AL:  "Transcriptional regulation of the human  carboxyl ester lipase gene in THP-1  monocytes: An E-box required for  activation binds upstream stimulatory  factors 1 and 2"  BIOCHEMICAL JOURNAL,  vol. 365, no. 2,  15 July 2002 (2002-07-15), pages 481-488,  XP002342784  ISSN: 0264-6021  the whole document  abstract  page 487, left-hand column, line 12 -  right-hand column, line 23</p>	1-9
A	<p>HUI DAVID Y ET AL: "Carboxyl ester  lipase: Structure-function relationship  and physiological role in lipoprotein  metabolism and atherosclerosis."  JOURNAL OF LIPID RESEARCH,  vol. 43, no. 12, December 2002 (2002-12),  pages 2017-2030, XP002342785  ISSN: 0022-2275  cited in the application  the whole document  abstract</p>	1-9
A	<p>BRUNEAU NADINE ET AL: "Lectin-like Ox-LDL  receptor is expressed in human INT-407  intestinal cells: Involvement in the  transcytosis of pancreatic bile  salt-dependent lipase."  MOLECULAR BIOLOGY OF THE CELL,  vol. 14, no. 7, July 2003 (2003-07), pages  2861-2875, XP002342786  ISSN: 1059-1524  cited in the application  the whole document</p>	1-9
A	<p>FALT HELEN ET AL: "Do human bile salt  stimulated lipase and colipase-dependent  pancreatic lipase share a common  heparin-containing receptor?"  ARCHIVES OF BIOCHEMISTRY AND BIOPHYSICS,  vol. 386, no. 2,  15 February 2001 (2001-02-15), pages  188-194, XP002342787  ISSN: 0003-9861  cited in the application  the whole document</p>	1-9

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C.(Continuation) DOCUMENTS CONSIDERED TO BE RELEVANT		
Category *	Citation of document, with indication, where appropriate, of the relevant passages	Relevant to claim No.
A	<p>CAILLOL NATHALIE ET AL: "Pancreatic bile salt-dependent lipase activity in serum of normolipidemic patients"</p> <p>LIPIDS, vol. 32, no. 11, November 1997 (1997-11), pages 1147-1153, XP008052126 ISSN: 0024-4201 cited in the application the whole document</p> <p>-----</p>	1-9

# INTERNATIONAL SEARCH REPORT

national application No.  
PCT/IB2005/051095

## Box II Observations where certain claims were found unsearchable (Continuation of item 2 of first sheet)

This International Search Report has not been established in respect of certain claims under Article 17(2)(a) for the following reasons:

1. ☒ Claims Nos.: 1-7, 9 (in part)  
because they relate to subject matter not required to be searched by this Authority, namely:  
Although claims 1-7 and 9 are directed to a method involving treatment of the human/animal body, the search has been carried out and based on the alleged effects of the compound/composition.
2. ☐ Claims Nos.:  
because they relate to parts of the International Application that do not comply with the prescribed requirements to such an extent that no meaningful International Search can be carried out, specifically:
3. ☐ Claims Nos.:  
because they are dependent claims and are not drafted in accordance with the second and third sentences of Rule 6.4(a).

## Box III Observations where unity of invention is lacking (Continuation of item 3 of first sheet)

This International Searching Authority found multiple inventions in this international application, as follows:

1. ☐ As all required additional search fees were timely paid by the applicant, this International Search Report covers all searchable claims.
2. ☐ As all searchable claims could be searched without effort justifying an additional fee, this Authority did not invite payment of any additional fee.
3. ☐ As only some of the required additional search fees were timely paid by the applicant, this International Search Report covers only those claims for which fees were paid, specifically claims Nos.:
4. ☐ No required additional search fees were timely paid by the applicant. Consequently, this International Search Report is restricted to the invention first mentioned in the claims; it is covered by claims Nos.:

Remark on Protest

- ☐ The additional search fees were accompanied by the applicant's protest.
- ☐ No protest accompanied the payment of additional search fees.

Patent document cited in search report	Publication date	Patent family member(s)	Publication date
EP 0640620 A	01-03-1995	AU 6610194 A CA 2127230 A1 JP 7165794 A	12-01-1995 02-01-1995 27-06-1995
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